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(54) Title: **ANTIMICROBIAL COMPOSITIONS FORMULATED FOR USE IN COLD TEMPERATURE CONDITIONS AND  
METHODS OF USE THEREOF**

(57) Abstract: Antimicrobial compositions containing a carboxylic acid, for example, a fatty acid, and a carrier medium including a freezing point depressant are disclosed. The compositions can be formulated for use as a teat dip, for use on milk producing animals. In one particularly advantageous embodiment, a composition is formulated as a teat dip and includes suitable emollients, skin conditioners and lubricants.

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## ANTIMICROBIAL COMPOSITIONS FORMULATED FOR USE IN COLD TEMPERATURE CONDITIONS AND METHODS OF USE THEREOF

5

### Field of the Invention

The present invention is directed to antimicrobial compositions having specialized benefit of skin protection and conditioning for the control of mastitis in milk producing animals that is especially useful in cold temperature environments and a method of treating for mastitis in cold temperature conditions. Furthermore, in  
10 some embodiments, the invention provides mastitis treatment compositions and formulations having specialized benefit of skin protection and conditioning on the teats of an animal that are already in poor health. In one preferred embodiment, the invention provides teat dip formulations comprising a carboxylic acid, typically a fatty acid, and a carrier medium including a freezing point depressing component.

15

### Background of the Invention

Mastitis is an inflammation of the mammary gland. Bovine mastitis is the most common and most costly disease affecting dairy herds. Some estimates suggest at least half of the dairy animal population have some degree or form of  
20 mastitis. This condition results in lowered milk yield and reduced milk quality. Economic loss to mastitis in the U.S. is estimated at about \$1.8 billion or approximately 10% of total milk sales with about two-thirds of this loss due to reduced milk production from infected cows. In dairy cattle, mastitis typically results from microorganisms; usually bacteria, that invade the udder, multiply in the  
25 milk producing tissues, and synthesize toxins, a by-product of bacterial metabolism.

The characteristic features of inflammation are swelling, heat, redness, pain and disturbed function.

While the animal immune system can fight intramammary infections, many chronic infections remain sub-clinical (asymptomatic) and undetected unless  
5 diagnosed by laboratory testing. Sub-clinical mastitis can result in a reservoir of microorganisms, which can lead to the infection of other animals within the herd. More than 80 species of microorganisms have been identified as causal agents, although approximately 95% of mastitis is caused by four pathogens:  
*Staphylococcus aureus*, *Streptococcus agalactiae*, *Streptococcus dysagalactiae*, and  
10 *Streptococcus uberis*. Mastitis causing pathogens fall into two categories namely contagious and environmental. Contagious bacteria, such as *Streptococcus agalactiae* and *Staphylococcus aureus*, primarily colonize host tissue sites such as mammary glands, teat canals, teat skin lesions etc. and are spread from one infected cow to another during the milking process. Environmental bacteria, often  
15 streptococci, enterococci and coliform organisms, are commonly present within the cow's surroundings from sources such as cow feces, soil, plant material, bedding or water, and infect by casual opportunistic contact with an animal. This distinction, although not exclusive, is of practical importance because different dairy herd maintenance measures are needed for the different groups of microorganisms. In all  
20 bovine mastitis cases, whatever the causal microorganism, the route of transmission of the invading pathogen into the inner gland of the udder is through the teat orifice and teat canal.

Management of dairy herds focuses attention on treatment of both established mastitis and on prevention of new intramammary infections. Therapy

and hygiene are two fundamental components of an effective mastitis control program. Each is applied in concert, and each operates independently. The primary effect of therapy is to eliminate established infections, whereas, hygiene reduces the incidence of new infection by interrupting transmission vectors. A non-exhaustive  
5 list of ancillary factors which may be employed for the elimination and prevention of mastitis, include, post-lactation antibiotic infusion into the udder (e.g., dry cow treatment); and, post-milking teat antisepsis or "teat dipping" during lactation.

Researchers agree, and an abundance of published evidence supports the concept, that dipping teats into an effective antimicrobial solution immediately after  
10 each milking is the single most effective procedure for decreasing new intramammary infections in lactating cows. Between 1955 to 1970, Dodd and co-workers conducted extensive epidemiologic investigations in commercial dairy herds (F. K. Neave, F. H. Dodd, and R. G. Kingwell, 1966, "A Method of Controlling Udder Disease", Vet. Rec. 78:521; F. K. Neave, F. H. Dodd, R.G.  
15 Kingwell and D. R. Westgarth, 1969, "Control of Mastitis in the Dairy Herd by Hygiene and Management", J. Dairy Sci. 52:696; F. H. Dodd, D. R. Westgarth, F. K. Neave and R. G. Kingwill, 1969, "Mastitis - The Strategy of Control", J. Dairy Sci. 52:689; and F. H. Dodd, and F. K. Neave, 1970, "Mastitis Control", Proceedings, Nat'l. Inst. Res. Dairying, pp. 21-60). From this work, they developed  
20 the conceptual basis for modern mastitis control methods of which teat dipping is an integral component. The efficacy and value of teat dipping has since been confirmed in dozens of field trials, and it is now accepted that an effective teat dip can reduce the incidence of new intramammary infections at least 50% and often up to 90%.

To reduce mastitis, commercial teat dips have been developed containing a variety of antimicrobial agents including iodophors, quaternary ammonium compounds, chlorhexidine salts, chlorine release compounds (e.g. alkali hypochlorites), oxidizing compounds (e.g. hydrogen peroxide, peracids), protonated carboxylic acids (e.g. octanoic, nonanoic, decanoic, acids), acid anionics (e.g. alkylaryl sulfonic acids), and chlorine dioxide (from chlorite). These agents, which have varying degrees of effectiveness, limit the transmission of mastitis by reducing pathogen populations on the teat. Teat dips, can also be divided into two broad classifications. The Class I type are antimicrobial and are applied to kill microorganisms already present in the teat canal or on the surface of the teat skin. By design, their microbiological effect is immediate and they primarily target the contagious organisms that are vectored between animals during the pre-milking, milking and post-milking process. The Class II type teat dip, often referred to as a "teat sealer," is a film-forming or coating composition which may or may not be antimicrobial; and, functions by developing a residual protective barrier on the teat thus providing prophylaxis by sealing the teat orifice from environmental contamination. The film, which forms on the surface of the teat, serves as a physical barrier through which mastitis causing pathogens cannot penetrate during the intermilking period.

General disclosures of teat dip technology are shown in, for example, "Current Concepts of Bovine Mastitis." 1996, Fourth Ed. National Mastitis Council, Madison WI.; P. A. Murdough and J. W. Pankey, 1993. "Evaluation of 57 Teat Sanitizers Using Excised Cow Teats", J. Dairy Sci. 76:2033-2038; J. W. Pankey et al., 1984, "Uptake on Post-milking Teat Antiseptics", J. Dairy Sci. 67:1336-1353; R.

- J. Farnsworth, 1980, "Role of Teat Dips in Mastitis Control", J. Am. Vet. Med. Assoc. 76:1116-1118; W. N. Philpot, 1979, "Control of Mastitis by Hygiene and Therapy", J. Dairy Sci. 62:168-176; W. N. Philpot and J. W. Pankey, 1978, "Hygiene in the Prevention of Udder Infections V. Efficacy of Teat Dips Under  
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10 M. Buckalew, 1972, "Evaluation of a Hygiene and Dry Period Therapy Program for Mastitis Control", J. Dairy Sci. 55:1683-1691; W. D. Schultze and J. W. Smith, 1972, "Effectiveness of Postmilking Teat Dips", J. Dairy Sci. 55:426-431; D. P. Wesen and L. H. Schultz, 1970, "Effectiveness of a Post-Milking Teat Dip in Preventing New Udder Infections", J. Dairy Sci. 53:1391-1403; and British Pat. No.  
15 1,144,637 (Kelco Chemicals Ltd.), published March 5, 1969.

Typical disclosures of intermilking or protective (barrier-type) film-forming teat dips or teat "sealers" can be found in Akers et. al., U. S. Pat. No. 3,066,071, issued November 27, 1962; Kraus, U.S. Pat. No. 3,222,252, issued December 7, 1965 (but, see Philpot et. al., J. Dairy Science 58:205-216); Coughman and Brown,  
20 U.S. Pat. No. 3,993,777, issued November 23, 1976; Pugliese, U. S. Pat. No. 4,049,830, issued September 20, 1977; and Andrews et al., U. S. Pat. No. 4,113,854, issued September 12, 1978. One disadvantage of many such film-forming agents is their tendency to form a "hard" film, which is tenacious and often difficult to remove.

There is a growing acceptance among academics, veterinarians and dairy herd management that proactive maintenance of teat health and skin condition is an integral part of a complete program for the prevention, control and remedial correction of mastitis in milk producing animals. A growing number of publications support this (see for example, M.D. McKinzie and T.C. Hemling, 1995, "The Effect of Teat Skin Condition on Milk Yield and Milkout Time", National Mastitis Council Proceedings, pp 160-163; L.K Fox, 1992, "Colonization of Staphylococcus Aureus On Chapped Skin: Effect of Iodine and Chlorhexidine Postmilking Teat Disinfectants", J. Dairy Sci. 75:66; and JJ Goldbert et al., 1994, "Evaluation of Teat Conditioning Qualities of Postmilking Teat Dips", National Mastitis Council technology transfer Session, Orlando, FL. The assertion is that a healthy milk delivery organ can naturally retard and more readily withstand the adverse affects of infection.

Although many teat dip products are available, and there remains a continuing need for new and effective teat dip compositions having immediate and long lasting antimicrobial effect against a wide spectrum of mastitis causing organisms; there is need for such antimicrobial compositions which additionally provide superior skin conditioning and health maintenance functions.

Additionally, many teat dip products are not conducive for use in cold environment conditions. For example, many teat dip compositions include a large amount of water, and can freeze in cold environments. Such compositions can freeze on the teats, and cause irritation, chapping and frostbite, and such conditions can enhance the incidence of mastitis caused by opportunistic invasive microorganisms.

Thus, although many teat dip products are available, there is a continuing need for new and effective teat dip compositions having immediate and long lasting antimicrobial effect against a wide spectrum of mastitis causing organisms that are formulated for use in cold environments.

5

### Summary of the Invention

The present invention is directed to novel antimicrobial compositions, which can be used as a teat dip for milk producing animals. At least some embodiments of this invention provide both superior antimicrobial protection against mastitis causing  
10 organisms and skin conditioning for maintenance of healthy teats; and in addition provide protection against the adverse affects of frigid weather, one of the major causes of irritation, chapping and frostbite, which enhance the incidence of mastitis caused by opportunistic invasive microorganisms. Still further benefit is accomplished by application of at least some preferred compositions of this  
15 invention on teats which are already in poor health condition, for example by application on so-termed "hospital pen animals", to facilitate faster repair and the healing process.

It will be noted that at several places throughout the Specification, guidance is provided through lists of examples. In each instance, the recited lists serve only as  
20 representative groups. It is not meant, however, that the lists are exclusive.

In general, an antimicrobial composition of the invention comprises, a C<sub>6</sub>-C<sub>12</sub> fatty acid and a carrier medium including a freezing point depressant component. In preferred embodiments, the fatty acid can be a C<sub>7</sub>-C<sub>9</sub> fatty acid. A particularly preferred fatty acid is heptanoic acid. In some preferred embodiments, the freezing



point depressant component makes up greater than 60% by wt. of the composition. Particularly preferred freezing point depressant components are selected from the group consisting of propylene glycol, glycerine, and mixtures thereof.

In preferred embodiments of a teat dip composition, the teat dip formulation  
5 also includes significantly high concentrations of one or more occlusive agents which provide a protective barrier on the teat surface against irritating physical abrasion as from the mechanical action of the milking equipment or tissue damaging or disruptive environmental conditions such as frigid temperatures, wind chill, dehydration, windburn and sunburn. These agents may be, as example, but not  
10 meant to be inclusive: surfactants, emollients, lubricants, humectants, moisturizers, solvents etc. or mixtures thereof having the purpose of generally protecting and conditioning the teat skin surfaces thereby promoting healthy milk producing animals. In some preferred embodiments, some of the freezing point depressant components have a dual function as a occlusive agent as well as a freezing point  
15 depressant component.

A teat dip composition can also include a rheology modifier, a film-forming agent or admixture, a buffer system, a hydrotrope or coupler, or solvent, an emollient, skin conditioner or a lubricant, surfactant, color marker, fragrance, anti-irritants and healing agents, antioxidants, UV absorbers, vitamins or admixtures thereof.

20 In general, the invention also includes a method of controlling mastitis in milk producing animals comprising applying the composition to a teat of an animal. The composition of the invention is particularly beneficial when applied in colder environmental temperatures, for example in temperatures of below 40°F.

### **Detailed Description of the Invention**

The present invention is directed to antimicrobial compositions. In general, the compositions include a fatty acid, typically a C<sub>6</sub>-C<sub>12</sub> fatty acid and a carrier medium including a freezing point depressant component. As used herein, "freezing point depressant component" means a constituent or mixture of constituents within the composition that act to reduce the freezing point of the composition below that of the composition without that constituent. Typically, the freezing point depressant component acts to reduce the freezing point of the composition below the freezing point of water. The compositions of the invention can be advantageously used as a teat dip for control of mastitis in milk producing animals. The compositions of the invention are formulated for use in cold temperature environments, for example in environments having a temperature of 40°F or below. However, formulating of the invention can also be used in warm environment conditions. Some embodiments are particularly useful to facilitate faster repair of teats that are already in poor health regardless of the temperature. For example, some embodiments are particularly beneficial on chapped or irritated teats.

### **Components Used in a Teat Dip Embodiment of the Invention**

The compositions of the invention can be formulated as a teat dip for mastitis prevention or control. According to this embodiment, the compositions can comprise a carboxylic acid, such as a fatty acid in a carrier medium comprising a freezing point depressant component. Preferably, the fatty acid comprises in the range of 0.01 to 5% by wt. of the total composition, and the freezing point depressant component comprises greater than 60% by wt. of the total composition.

Teat dip compositions may optionally also include, a rheology modifier or admixture, a film-forming agent or admixture, a buffer system, a hydrotrope, a coupler or admixtures thereof, a surfactant or surfactant admixture, an emollient, a skin-conditioner or a lubricant or admixtures thereof, one or more adjuvants, etc.

- 5 The preferred compositions of the invention comprise ingredients, which are generally regarded as safe, and are not of themselves or in admixture incompatible with milk or milk by-products. Ingredients may also be selected which are cooperative in their combined effects whether incorporated for physical integrity of the formulation or to facilitate healing and overall health of the teat. The carrier  
10 functions to dilute the active ingredients and facilitate application to the intended surface.

- The compositions of the invention can be provided as a ready-to-use formulation or as concentrates, which are diluted prior to use. Hence, throughout this disclosure, reference will be made to "working compositions" which are the  
15 compositions, which are actually used as a teat dip. Thus, a working composition includes, for example, ready-to-use compositions as well as concentrates, which have been diluted for use in a particular application. Methods for preparing concentrates based on the disclosure herein of working compositions are within the knowledge of one of skill in the art.

- 20 In preferred embodiments, a teat dip composition of the invention provides a protective soft barrier over the teat. A "soft barrier" provides a self annealing barrier which can flow to re-cover areas of the teat from which the dip may have been removed when the animal lays down, walks through the pasture or is subject to some other event which causes inadvertent removal of the dip. Advantageously, however,

a herein disclosed soft barrier can also be readily removed from the teat using routine washing procedures prior to milking without congealing, pilling or leaving some other undesired residue on the teat. The soft barrier is provided and rheology modifiers described further below. Thus, unlike prior film-forming barriers, the protective soft barrier provided herein is a soft, non-peeling barrier that can undergo plastic deformation, for self-repair, without breaking or cracking.

#### **Carrier Medium Including Freezing Point Depressant Component**

The carrier medium of a composition of the invention can generally be an aqueous medium such as water, or an organic liquid such as an oil, a surfactant, an alcohol or polyol, an ester, an ether, or an organic or aqueous mixture of any of these. The carrier medium includes high concentrations of non-aqueous, preferably water miscible liquids or soluble solids that act as a freezing point depressant component for the composition. Preferred freezing point depressant components also have some degree of skin conditioning properties, and provide protection against the adverse affects of frigid weather. Examples of preferred freezing point depressant components include polyols, and mixtures thereof. Preferred polyols include propylene glycol; glycerin; sorbitol; and homologs, homopolymers and derivatives thereof and mixtures thereof. Preferably, the freezing point depressant component of the carrier medium make up greater than 60%, in some embodiments, greater than 65%, in some other embodiments, greater than 70%, and in yet other embodiments, greater than 75% by weight of the total composition.

The freezing point depressant components within the composition have the desired effect of reducing the freezing point of the composition below that of water.

In some embodiments, the freezing point of the composition is below 25°F, in some embodiments, below 15°F, in some embodiments, below 10°F, in some embodiments below 0°F, in some embodiments below -10°F, and in yet other embodiments, below -20°F. By providing the composition with a low freezing point, the freezing point depressants allow the composition to be used in cold temperature environments, and help to prevent chapping, irritation, windburn, frostbite, and other such conditions on the teats of an animal associated with cold environments. Further, the compositions are less likely to freeze and can be applied in colder temperatures than many teat dips.

10

### Fatty Acid

The compositions of the invention also include a carboxylic fatty acid component, which provides the antimicrobial activity. For optimal antimicrobial activity, the pH of the composition is preferably at or below the pKa of the fatty acid. Thus, in preferred compositions, the relationship of pH to pKa provides antimicrobial activity through carboxylic fatty acids, which are substantially protonated. Preferred compositions of the invention have a pH in the range of about 3.5 to about 6.0, typically about 4.0 to about 5.0 and, in one preferred embodiment, about 4.5. Although a wider range of pH is possible, typically, below pH 3.5 undesirable skin irritation may occur and above pH 6.0 dissociation or conversion to the ionized form may reduce the antimicrobial efficacy of the fatty acid.

20

Fatty acids suitable for a composition of the present invention include C<sub>6</sub>-C<sub>12</sub> fatty acids. Preferred fatty acids have a chain length from about C<sub>7</sub>-C<sub>9</sub>. One particularly preferred fatty acid is heptanoic acid that has seven carbon atoms,

including the carboxyl group, and has a pKa of 4.4. In addition to its preferred water solubility, heptanoic acid is not significantly irritating to the tissues.

In general, the fatty acid component of the invention can be present at about 0.01% to about 5.0% of the total weight of the working composition. For example, 5 when heptanoic acid is the fatty acid, it may be present at about 0.01% to about 5% of the total weight of the working composition. In a working composition having a pH of about 4.5, heptanoic acid may be present at about 0.1% to about 5% of the total working solution, preferably at about 0.5% to about 2.0%.

The antimicrobial components of the composition can be mixed in the carrier 10 medium and include buffers, surface active agents and/or couplers to provide a pH and solubility suitable for efficient bactericidal effect with low or no irritation to the tissues of the teat. The buffer system is present to prevent the likelihood of pH drift under typical use conditions. In general, the buffer system can include any weak acid and its conjugate base. Preferred bases used to adjust the pH of the 15 compositions include hydroxides of the alkaline earth metals, for example NaOH, KOH, LiOH, etc.

Maintenance of the pH of compositions described in this invention is preferred to minimize undesirable chemical changes, which may inhibit the microbiological efficacy of the antimicrobial components or cause toxic or irritating 20 effect upon the teat. Any compatible organic or inorganic material or mixture of materials which has the desired effect of maintaining the composition pH within prescribed ranges can be utilized as the buffering agent or system in the invention. Factors which may cause undesirable pH shifts include the presence of naturally occurring chemicals brought into the composition, after application onto the teat, by

skin exudations, milk or environmental soils; and, pH drifting which sometimes accompanies chemical equilibriums established within compositions as ingredients are changed or concentrations varied, for example, concentration changes which can occur as a teat dip dries on the teat.

5           In general, the pH of bovine mastitis control treatments can vary from a low of about pH 2.5 to a maximum of approximately 10.5 depending primarily upon the choice of antimicrobial agent being incorporated in the composition. Therefore the buffering agent or system is chosen accordingly. Most common commercially-available weak inorganic and organic acids can be used in the  
10   invention. Preferred weak inorganic acids include phosphoric acid and sulfamic acid. Useful weak organic acids include acetic acid, hydroxyacetic acid, citric acid, tartaric acid, lactic acid, glycolic acid, adipic acid, succinic acid, propionic acid, malic acid, alkane sulfonic acids, cycloalkane sulfonic acids, etc. Mixtures of organic and inorganic acids can also be used. One typical and preferred buffer  
15   system is citric acid and its alkali metal salt.

Solubilizing agents called hydrotropes or couplers or solvents may be generally used in compositions of the invention to maintain physical single-phase integrity and storage stability. To this end, any number of ingredients known to those skilled in the formulation art may be employed, such as monofunctional and  
20   polyfunctional alcohols. These preferably contain from about 1 to about 6 carbon atoms and from 1 to about 6 hydroxy groups. Examples include ethanol, isopropanol, n-propanol, 1, 2-propanediol, 1, 2-butanediol, 2-methyl-2, 4-pentanediol, mannitol and glucose. Also useful are the higher glycols, polyglycols, polyoxides, glycol ethers and propylene glycol ethers. Additional useful

hydrotropes include the free acids and alkali metal salts of sulfonated alkylaryls such as toluene, xylene, cumene and phenol or phenol ether or diphenyl ether sulfonates; alkyl and dialkyl naphthalene sulfonates and alkoxylated derivatives.

#### Additional Components

5           A composition of the invention may also contain one or more rheology modifiers, to enhance viscosity, or thicken the composition to facilitate adherence of a dip to the teat. Adherence enables the composition to remain in contact with transient and resident pathogenic bacteria for longer periods of time, promoting microbiological efficacy and resisting waste because of excessive dripping. The  
10   rheology modifier may be a film former or act cooperatively with a film-forming agent to form a barrier that provides additional protection. However, in preferred embodiments, a teat dip composition of the invention provides a soft barrier, rather than a well-defined film.

          Water soluble or water dispersible rheology modifiers that are useful can be  
15   classified as inorganic or organic. The organic thickeners can further be divided into natural and synthetic polymers with the latter still further subdivided into synthetic natural-based and synthetic petroleum-based.

          Inorganic thickeners are generally compounds such as colloidal magnesium aluminum silicate (VEEGUM®), colloidal clays (Bentonites), or silicas (CAB-O-  
20   SILS®) which have been fumed or precipitated to create particles with large surface to size ratios. Suitable natural hydrogel thickeners are primarily vegetable derived exudates. For example, tragacanth, karaya, and acacia gums; and extractives such as carrageenan, locust bean gum, guar gum and pectin; or, pure culture fermentation products such as xanthan gum. Chemically, all of these materials are salts of



complex anionic polysaccharides. Synthetic natural-based thickeners having application are cellulosic derivatives wherein the free hydroxyl groups on the linear anhydro-glucose polymers have been etherified or esterified to give a family of substances, which dissolve in water and give viscous solutions. This group of materials includes the alkyl and hydroxylalkylcelluloses, specifically methylcellulose, hydroxyethylmethylcellulose, hydroxypropylmethylcellulose, hydroxybutylmethylcellulose, hydroxyethylcellulose, ethylhydroxyethylcellulose, hydroxypropylcellulose, and carboxymethylcellulose. Synthetic petroleum-based water soluble polymers are prepared by direct polymerization of suitable monomers of which polyvinylpyrrolidone, polyvinylmethylether, polyacrylic acid and polymethacrylic acid, polyacrylamide, polyethylene oxide, and polyethyleneimine are representative.

A preferred rheology modifier is polyvinylpyrrolidone, K-90 manufactured by International Specialty Products. This rheology modifier is particularly advantageous in that it is very water soluble, compatible in compositions containing high formula percentages of organic materials and is believed to retard skin irritation.

Suitable surfactants or surfactant admixtures can be selected from compatible water-soluble or water dispersible nonionic, or anionic surface-active agents; or mixtures of each or both types. Non-ionic surfactants useful in the invention are generally characterized by the presence of an organic hydrophobic group and an organic hydrophilic group and are typically produced by the condensation of an organic aliphatic, alkyl aromatic or polyoxyalkylene hydrophobic compound with a hydrophilic alkaline oxide moiety which in common practice is ethylene oxide or a

polyhydration product thereof, polyethylene glycol. Practically any hydrophobic compound having a hydroxyl, carboxyl, amino, or amido group with a reactive hydrogen atom can be condensed with ethylene oxide, or its polyhydration adducts, or its mixtures with alkoxylenes such as propylene oxide to form a nonionic surface-active agent. The length of the hydrophilic polyoxyalkylene moiety which is condensed with any particular hydrophobic compound can be readily adjusted to yield a water dispersible or water soluble compound having the desired degree of balance between hydrophilic and hydrophobic properties.

Useful nonionic surfactants in the present invention include:

- 10 Block polyoxypropylene-polyoxyethylene polymeric compounds based upon propylene glycol, ethylene glycol, glycerol and trimethylolpropane as the initiator reactive hydrogen compound. Examples of polymeric compounds made from a sequential propoxylation and ethoxylation of initiator are commercially available under the trade name PLURONIC<sup>®</sup> manufactured by BASF Corp. PLURONIC<sup>®</sup>
- 15 compounds are difunctional (two reactive hydrogens) compounds formed by condensing ethylene oxide with a hydrophobic base formed by the addition of propylene oxide to two hydroxyl groups of propylene glycol. This hydrophobic portion of the molecule weighs from about 1,000 to about 4,000. Ethylene oxide is then added to sandwich this hydrophobe between hydrophilic groups, controlled by
- 20 length to constitute from about 10% by weight to about 80% by weight of the final molecule.

Likewise useful nonionic surfactants include condensation products of one mole of a saturated or unsaturated, straight or branched chain alcohol having from about 6 to about 24 carbon atoms with from about 3 to about 50 moles of ethylene

oxide. The alcohol moiety can consist of mixtures of alcohols in the above delineated carbon range or it can consist of an alcohol having a specific number of carbon atoms within this range. Examples of like commercial surfactant are available under the trade name NEODOL<sup>®</sup> manufactured by Shell Chemical Co. and

5 ALFONIC<sup>®</sup> manufactured by Vista Chemical Co.

Condensation products of one mole of saturated or unsaturated, straight or branched chain carboxylic acid having from about 8 to about 18 carbon atoms with from about 6 to about 50 moles of ethylene oxide. The acid moiety can consist of mixtures of acids in the above delineated carbon atoms range or it can consist of an

10 acid having a specific number of carbon atoms within the range. Examples of commercial compounds of this chemistry are available on the market under the trade name NOPALCOL<sup>®</sup> manufactured by Henkel Corporation and LIPOPEG<sup>®</sup> manufactured by Lipo Chemicals, Inc. In addition to ethoxylated carboxylic acids, other alkanoic acid esters formed by reaction with glycerides, glycerin, and

15 polyhydric (saccharide or sorbitan/sorbitol) alcohols have application in this invention. All of these ester moieties have one or more reactive hydrogen sites on their molecule which can undergo further acylation or ethylene oxide (alkoxide) addition to control the hydrophilicity of these substances.

Also useful nonionic surfactants include the condensation products of one

20 mole of alkyl phenol wherein the alkyl constituent contains from about 8 to about 18 carbon atoms with from about 3 to about 50 moles of ethylene oxide. The alkyl group can, for example, be represented by diisobutylene, di-amyl, polymerized propylene, isoctyl, nonyl, and di-nonyl. Examples of commercial compounds of this chemistry are available on the market under the trade name IGEPAL<sup>®</sup> manufactured

by Rhone-Poulenc and TRITON<sup>®</sup> manufactured by Union Carbide. The surfactants used in the present compositions are also selected to improve solubility for removal of the composition from the teat prior to milking.

Teat dip compositions of the present invention can also include an emollient,  
5 moisturizer, humectant or re-fatting agent to lubricate, condition and generally reduce irritation and promote the healing of the teat surface of which may result either from the antimicrobial component, from the mechanical action of the milking machine or from environmental conditions such as frigid temperatures wind chill, dehydration, abrasion, windburn and sunburn. Any water miscible, soluble or  
10 dispersible skin-conditioning agent may be used in this invention. Compositions such as polyhydric alcohols are useful in the invention including glycerin, sorbitol, mannitol, and propylene glycol and its homopolymers; fatty acid esters of simple monohydric alcohols including isopropyl palmitate or isopropyl myristate and similar esters; polyol esters of fatty acids; and, ethoxylated lanolins, vegetable oils,  
15 and similar natural sourced derivatives such as aloe. Preferred emollients to be used in the invention include glycerin, and propylene glycol and lanolin. It should be noted that preferably the freezing point depressant component in the carrier medium also act as emollient/moisturizer/humectants. For example, in some preferable embodiments, propylene glycol and glycerin are present in high concentrations in  
20 the composition, and act as both freezing point depressants and emollients.

The compositions of the invention may also optionally include medicaments, for example sunscreens such as paraamino benzoic acid and healing agents such as allantoin or tocopheryl acetate or  $\alpha$ -hydroxy carboxylic acids or urea to provide curative action and stimulation of formation of new tissue; preservatives such as

methy1 paraben, propyl paraben, sorbic and benzoic acids or salts thereof to retard bacterial growth and prolong shelf life; antioxidants such as BHT (butylated hydroxytoluene), BHA (butylated hydroxyanisole), TBHQ (tert-butylhydroquinone), or propyl gallate to retard oxidative or hydrolytic degradation; sequestering agents  
5 such as aminopolyacetates, polyphosphonates, aminopolyphosphonates, polycarboxylates, and condensed phosphates; dispersants or suspending agents having polyelectrolytic character such as polyacrylate and similar polycarboxylates of homopolymeric or copolymeric structure; and manufacturing processing agents, for example defoam additives employed to facilitate blending and mixing.

10 The compositions can also include, dyes, pigments, marking agents, or other such components, as is generally known.

#### **Preferred Methods of Using the Compositions**

The compositions of the invention are useful in a method for controlling  
15 mastitis in milk producing animals. Generally, the method includes applying an antimicrobial composition embodying the invention a teat of an animal. Due to the freezing point depressant component within the composition, the compositions of the invention are particularly useful for methods wherein the antimicrobial composition is applied in environmental temperatures of below 40°F or is applied to  
20 the teat of an animal that will be exposed to environmental temperatures of below 40°F within 12 hours of the application. In some embodiments of such methods, the environmental temperatures can be below 30°F, below 20°F, or below 10°F.

Although the compositions of the invention are generally formulated for use in colder environmental temperatures, as discussed above, at least some embodiments

of the compositions can be used in methods of treating teats on a milk producing animal which are already in poor health condition, for example by application on so-called "hospital pen animals", to facilitate faster repair and the healing process.

The following examples are provided to further describe certain  
5 advantageous compositions according to the invention. The Examples, however, are not intended to limit the scope of the compositions within the spirit and scope of the invention.

### Examples

#### 10 Example 1 Formulation of Teat Dip Composition

The present Example provides one procedure for preparing a working composition of a teat dip composition according to one embodiment of the invention. This procedure can be used regardless of the total weight of the composition formulated. Thus, while a particular weight percentage of a component  
15 may vary among formulations, the procedure used for mixing the components is the same. It will be appreciated that other procedures can be used and are within the knowledge of one skilled in the art.

Deionized water is added to a stainless steel tank having a variable speed pitched blade turbine. The tank is agitated and the surfactant, sodium  
20 dioctylsulfosuccinate, is then added, with about 15 minutes of mixing and allowed to solubilize the surfactant. Polyvinylpyrrolidone K-90 is charged and mixed until completely solubilized. Preferably, the Polyvinylpyrrolidone is completely solubilized prior to addition of subsequent components.

Propylene glycol (technical), glycerin (e.g., 96% USP), ethoxylated lanolin, the fatty acid (e.g., heptanoic acid) and the fragrance, methyl solicylate are added and mixed for about 15 minutes to incorporate them into the mixture. A buffer, such as anhydrous granular citric acid is slowly added to the mixture and mixed until  
5 solubilized. Liquid 45% potassium hydroxide and blue dye are then added and the entire formula blended until the mixture appearance is uniform.

The preferred pH for a teat dip composition is about 3.5 to about 6. If the pH is less than the preferred range, a base such as potassium hydroxide can be added incrementally until the appropriate pH is achieved. If the pH is greater than the  
10 preferred range, an acid such as phosphoric acid can be added incrementally until the appropriate pH is obtained.

Tables 1-7 provide exemplary formulations for some teat dip compositions according to the invention.

Table 2: Teat Dip Formulations 2-9

ITEM	RAW MATERIAL	RM CODES	2	3	4	5	6	7	8	9
			WT %	WT %	WT %	WT %	WT %	WT %	WT %	WT %
1	DI Water	100032	22.650	22.503	23.347	22.498	23.347	23.350	23.355	23.400
2	Neodol 25-9	173534	0.100	0.100	0.100					
3	AEROSOL OT	171000				0.100	0.100	0.100	0.100	0.100
4	Propylene Glycol	164079	75.000	75.000	75.000	75.000	75.000	75.000	75.000	75.000
5	Heptanoic Acid, C-7	190454	1.500	1.500	1.500	1.500	1.500	1.500	1.500	1.500
6	Lactic Acid, 88%	810432	0.750	0.897		0.902				
7	Phosphoric Acid, 75%	121012			0.053		0.053	0.050	0.045	
		Total	100.000	100.000	100.000	100.000	100.000	100.000	100.000	100.000

Neodol 25-9: Shell Chemical Co., Houston, TX.



Table 3: Teat Dip Formulations 10-17

ITEM	RAW MATERIAL	RM CODES	10	11	12	13	14	15	16	17
			WT %	WT %	WT %	WT %	WT %	WT %	WT %	WT %
1	DI Water	100032	23.393	23.375	23.325	23.313	23.307	20.340	23.250	20.350
2	Neodol 25-9	173534								
3	AEROSOL OT	171000	0.100	0.100	0.100	0.100	0.100	0.020	0.100	0.100
4	PVP K-90	230076							0.100	
5	Rheothick 80 -11, 16%									3.000
6	Propylene Glycol	164079	75.000	75.000	75.000	75.000	75.000	73.000	75.000	75.000
7	Glycerine, 96%	164186						5.000		
8	Laneto 50	810762						0.100		
9	Heptanoic Acid, C-7	190454	1.500	1.500	1.500	1.500	1.500	1.500	1.500	1.500
10	Phosphoric Acid, 75%	121012				0.045	0.048	0.040	0.050	0.050
11	KOH, 45%	114041	0.007	0.025	0.075					
12	FD&C Yellow # 6	810739				0.042	0.045			
	Total		100.000	100.000	100.000	100.000	100.000	100.000	100.000	100.000

Rheothick 80-11: Henkel Corp, Chemicals Group Care Chemicals, Ambler, PA

### **Table 4: Teat Dip Formulations 18-25**

[illegible]

Table 5: Teat Dip Formulations 26-33

ITEM	RAW MATERIAL	RM CODES	26	27	28	29	30	31	32	33
			WT %	WT %	WT %	WT %	WT %	WT %	WT %	WT %
1	DI Water	100032	20.118	20.099	20.160	19.179	19.133	19.152	19.194	19.312
2	AEROSOL OT	171000	0.020	0.020	0.020	0.020	0.020	0.020	0.020	0.020
3	KLUCCEL HF	230151	0.100	0.100	0.100					
4	PVP K-90	230076				1.000	1.000	1.000	1.000	1.000
6	Propylene Glycol	164079	73.000	73.000	73.000	73.132	73.132	73.132	73.132	73.000
7	Glycerine, 96%	164186	5.000	5.000	5.000	5.000	5.000	5.000	5.000	5.000
8	Laneto 50	810762	0.100	0.100	0.100	0.100	0.100	0.100	0.100	0.100
9	Heptanoic Acid, C-7	190454	1.500	1.500	1.500	1.500	1.500	1.500	1.500	1.500
10	Phosphoric Acid, 75%	121012	0.012	0.012	0.012					
11	KOH, 45%	114041	0.008	0.008	0.008	0.004	0.004	0.004	0.004	0.003
12	Benzaldehyde	283028				0.050	0.050	0.050	0.050	
13	Vanillin	260607	0.100	0.100	0.100					
14	Methyl Salicylate	261016								0.050
15	FD&C Yellow # 5	271684	0.002	0.060			0.060	0.002		
16	FD&C Blue # 1	810184		0.001		0.015	0.001			0.015
17	FD&C Red # 3	274449	0.040					0.040		
			100.000	100.000	100.000	100.000	100.000	100.000	100.000	100.000
		Total								

Table 6: Teat Dip Formulations 34-41

ITEM	RAW MATERIAL	RM CODES	34	35	36	37	38	39	40	41
			WT %	WT %	WT %	WT %	WT %	WT %	WT %	WT %
1	DI Water	10032	19.267	19.306	19.283	20.161	19.309	20.207	19.335	18.695
2	AEROSOL OT	171000	0.020	0.020	0.020	0.020	0.020	0.020		0.020
3	KLUCEL HF	230151				0.100		0.100		
4	PVP K-90	230076	1.000	1.000	1.000		1.000		1.000	1.000
5	Propylene Glycol	164079	73.000	73.000	73.000	73.000	63.000	63.000	63.000	63.000
6	Glycerine, 96%	164186	5.000	5.000	5.000	5.000	15.000	15.000	15.000	15.000
7	Laneto 50	810762	0.100	0.100	0.100	0.100	0.100	0.100	0.100	0.100
8	Heptanoic Acid, C-7	190454	1.500	1.500	1.500	1.500	1.500	1.500	1.500	1.500
9	KOH, 45%	114041	0.003	0.003	0.003	0.008	0.006	0.008		0.135
10	Citric Acid, Anhyd.	124032								0.500
11	Methyl Salicylate	261016	0.050	0.050	0.050	0.050	0.050	0.050	0.050	0.050
12	FD&C Yellow # 5	271684	0.060	0.001		0.060				
13	FD&C Yellow # 6	810739			0.045					
14	FD&C Blue # 1	810184	0.001			0.001	0.015	0.015	0.015	
15	FD&C Red # 3	274449		0.020						
			100.000	100.000	100.000	100.000	100.000	100.000	100.000	100.000
		Total								

*Klucel HF: Hercules Inc., Aqualon Div., Wilmington, DE*

**Table 7: Teat Dip Formulations 42-47**

ITEM	RAW MATERIAL	RM CODES	42	43	44	45	46	47
			WT %	WT %	WT %	WT %	WT %	WT %
1	DI Water	100032	18.676	18.543	19.298	19.167	18.790	18.776
2	AEROSOL OT	171000	0.020	0.020	0.020	0.020	0.020	0.020
3	PVP K-90	230076	1.000	1.000	0.499	0.500	0.500	0.500
4	Propylene Glycol	164079	63.000	63.000	62.897	63.000	63.000	63.000
5	Glycerine, 96%	164186	15.000	15.000	14.975	15.000	15.000	15.000
6	Laneto 50	810762	0.100	0.100	0.100	0.100	0.100	0.100
7	Heptanoic Acid, C-7	190454	1.500	1.500	1.498	1.500	1.750	1.750
8	KOH, 45%	114041	0.139	0.272	0.150	0.148	0.290	0.289
9	Citric Acid, Anhyd.	124032	0.500	0.500	0.499	0.500	0.500	0.500
10	Methyl Salicylate	261016	0.050	0.050	0.050	0.050	0.050	0.050
11	FD&C Blue # 1	810184	0.015	0.015	0.015	0.015		0.015
		Total	100.000	100.000	100.000	100.000	100.000	100.000

**Example 2 Sanitizing Efficacy of Teat Dip KX-6128 against Eight Mastitis Causing Organisms**

The objective of the analysis was to determine the sanitizing efficacy of the teat dip composition 1 (from Table 1 above) against *Staphylococcus aureus* ATCC 6538, *Escherichia coli* ATCC 11229, *Klebsiella pneumoniae* ATCC 4352, *Streptococcus agalactiae* ATCC 27056, *Streptococcus dysgalactiae* ATCC 27957, *Streptococcus uberis* ATCC 27958, *Enterobacter aerogenes* ATCC 13048, and *Pseudomonas aeruginosa* ATCC 15442 at a 15 second exposure time with a 10% milk challenge.

**TEST METHOD:**

The method used in this example generally followed AOAC method 960.09, with the following deviations: the exposure contact time was reduced to 15 seconds from 30 seconds. Additionally, a 10% whole milk soil challenge was prepared by adding 10 mls of raw whole milk to 90 mls of test product and mixed, and from which 99 ml aliquot is used for the test (with 1 ml of specified inoculum).

**METHOD PARAMETERS:**

Test Substance Name	Concentration	mL of Test Substance	mL of Diluent
KX-6128	100%	90	10

Test Systems:	<i>Staphylococcus aureus</i>	ATCC 6538
	<i>Escherichia coli</i>	ATCC 11229
	<i>Klebsiella pneumoniae</i>	ATCC 4352
	<i>Enterobacter aerogenes</i>	ATCC 13048
	<i>Streptococcus agalactiae</i>	ATCC 27956
	<i>Streptococcus dysgalactiae</i>	ATCC 27957
	<i>Streptococcus uberis</i>	ATCC 27958
	<i>Pseudomonas aeruginosa</i>	ATCC 15442

Test Temperature: Ambient temperature

**Exposure Time:** 15 seconds

**Neutralizer:** Chambers

**Subculture Media:** Tryptone Glucose Extract Agar  
Brain Heart Infusion Agar (for *Streptococci*)

**Incubation:** 37°C for 48 hours

## RESULTS:

### Inoculum Numbers (CFU/mL)

Organism	Date	A	B	C	Average
<i>S. aureus</i> ATCC 6538	7/18/00	94 x 10 <sup>6</sup>	91 x 10 <sup>6</sup>	90 x 10 <sup>6</sup>	9.2 x 10 <sup>7</sup>
<i>E. coli</i> ATCC 11229	6/29/00	115 x 10 <sup>6</sup>	105 x 10 <sup>6</sup>	102 x 10 <sup>6</sup>	1.1 x 10 <sup>8</sup>
<i>E. aerogenes</i> ATCC 13048	6/29/00	70 x 10 <sup>6</sup>	66 x 10 <sup>6</sup>	64 x 10 <sup>6</sup>	6.7 x 10 <sup>7</sup>
<i>Klebsiella pneumoniae</i> ATCC 4352	7/18/00	168 x 10 <sup>6</sup>	176 x 10 <sup>6</sup>	126 x 10 <sup>6</sup>	1.6 x 10 <sup>8</sup>
<i>Streptococcus agalactiae</i> ATCC 27956	6/29/00	36 x 10 <sup>6</sup>	58 x 10 <sup>6</sup>	28 x 10 <sup>6</sup>	4.1 x 10 <sup>7</sup>
<i>Streptococcus dysgalactiae</i> ATCC 27957	7/18/00	74 x 10 <sup>6</sup>	59 x 10 <sup>6</sup>	44 x 10 <sup>6</sup>	5.9 x 10 <sup>7</sup>
<i>Streptococcus uberis</i> ATCC 27958	7/18/00	67 x 10 <sup>6</sup>	76 x 10 <sup>6</sup>	38 x 10 <sup>6</sup>	6.0 x 10 <sup>7</sup>
<i>Pseudomonas aeruginosa</i> ATCC 15442	7/18/00	84 x 10 <sup>6</sup>	163 x 10 <sup>6</sup>	134 x 10 <sup>6</sup>	1.3 x 10 <sup>8</sup>

**Test Results**

Test Substance	Test Organism	Survivors (CFU/mL)	Average Survivors (CFU/mL)	Log Reduction	Percent Reduction
KX-6128	<i>S. aureus</i>	<10, <10	<10	>6.96	>99.999
KX-6128	<i>E. coli</i>	<10, <10	<10	>7.04	>99.999
KX-6128	<i>E. aerogenes</i>	<10, <10	<10	>6.83	>99.999
KX-6128	<i>K. pneumoniae</i>	<10, <10	<10	>7.20	>99.999
KX-6128	<i>S. agalactiae</i>	<10, $2.0 \times 10^1$	$1.5 \times 10^1$	>6.61	>99.999
KX-6128	<i>S. dysgalactiae</i>	<10, <10	<10	>6.77	>99.999
KX-6128	<i>S. uberis</i>	<10, <10	<10	>6.78	>99.999
KX-6128	<i>P. aeruginosa</i>	<10, <10	<10	>7.11	>99.999

5

**CONCLUSIONS:**

Teat Dip Formulation 1 (from table 1) achieved a >99.999 percent reduction with a 15 second exposure time at ambient temperature against *Staphylococcus aureus* ATCC 6538, *Escherichia coli* ATCC 11229, *Enterobacter aerogenes* ATCC 13048, *Klebsiella pneumoniae* ATCC 4352, *Streptococcus agalactiae* ATCC 27956, *Streptococcus dysgalactiae* ATCC 27958, *Streptococcus uberis* ATCC 27958, and *Pseudomonas aeruginosa* ATCC 15442. This percent reduction was unexpectedly high.

15

From the foregoing detailed description and examples, it will be evident that modifications and variations can be made to the compositions and methods of the invention without departing from the spirit and scope of the invention. Therefore, it is intended that all modifications made to the invention without departing from the spirit and scope of the invention come within the scope of the appended claims.



**WHAT IS CLAIMED IS:**

1. An antimicrobial composition comprising:  
in the range of 0.01 to 5 wt. % of a C6-C12 fatty acid; and  
a carrier medium including a freezing point depressant component, wherein the  
5 freezing point depressant component makes up greater than 60 wt. % of the total  
composition.
2. The antimicrobial composition of claim 1, wherein the fatty acid is C7-C9.
- 10 3. The antimicrobial composition of claim 1, wherein the fatty acid is heptanoic  
acid.
4. The antimicrobial composition of claim 1, wherein the freezing point depressant  
component is selected from polyol or mixtures thereof.
- 15 5. The antimicrobial composition of claim 4, wherein the freezing point depressant  
component is selected from propylene glycol, glycerin, and mixtures thereof.
6. The antimicrobial composition of claim 5, wherein the freezing point depressant  
20 component is a mixture of propylene glycol and glycerin.
7. The antimicrobial composition of claim 1, wherein the composition has a  
freezing point of below 32°F.

8. The antimicrobial composition of claim 1, wherein the composition has a freezing point of below 20°F.
9. The antimicrobial composition of claim 1, wherein the composition has a  
5 freezing point of below 10°F.
10. The antimicrobial composition of claim 1, wherein the composition has a freezing point of below 0°F.
- 10 11. The antimicrobial composition of claim 1, wherein the composition has a freezing point of below -10°F.
12. The antimicrobial composition of claim 1, wherein the composition has a freezing point of below -20°F.
- 15 14. The antimicrobial composition of claim 1, wherein the freezing point depressant component makes up greater than 65 wt. % of the total composition.
15. The antimicrobial composition of claim 1, wherein the freezing point depressant  
20 component makes up greater than 70 wt. % of the total composition.
16. The antimicrobial composition of claim 1, wherein the freezing point depressant component makes up greater than 75 wt. % of the total composition.
- 25 17. An antimicrobial composition comprising:

in the range of 0.01 to 5 wt. % of heptanoic acid; and  
greater than 60 wt. % of a freezing point depressant component comprising  
propylene glycol and glycerin.

5 18. An antimicrobial composition comprising:

an antimicrobial component consisting essentially of heptanoic acid, and  
a carrier medium.

19. A method for controlling mastitis in milk producing animals, the method  
10 comprising:

applying an antimicrobial composition to a teat of an animal wherein the  
antimicrobial composition comprises in the range of 0.01 to 5 wt. % of a C6-C12 fatty  
acid and a carrier medium including a freezing point depressant component, wherein  
the freezing point depressant component comprises greater than 60 wt. % of the

15 composition.

20. The method of claim 19, wherein the antimicrobial composition is applied in  
environmental temperatures of below 40°F or is applied to the teat of an animal that  
will be exposed to environmental temperatures of below 40°F within 12 hours of the  
20 application.

21. The method of claim 20, wherein the environmental temperatures of below  
30°F.

22. The method of claim 20, wherein the environmental temperatures of below 20°F.

23. The method of claim 20, wherein the environmental temperatures of below  
5 10°F.

24. The method of claim 19, wherein the fatty acid is C7-C9.

25. The method of claim 19, wherein the fatty acid is heptanoic acid.

10

26. The method of claim 19, wherein the freezing point depressant component is selected from polyols or mixtures thereof.

27. The method of claim 26, wherein the freezing point depressant component is  
15 selected from the group consisting of propylene glycol, glycerin, and mixtures thereof.

28. The method of claim 27, wherein, wherein the freezing point depressant component is a mixture of propylene glycol and glycerin.

20 29. The method of claim 19, wherein the composition has a freezing point of below 32°F.

30. The method of claim 19, wherein the composition has a freezing point of below 20°F.

25

31. The method of claim 19, wherein the composition has a freezing point of below 10°F.
32. The method of claim 19, wherein the composition has a freezing point of below  
5 0°F.
33. The method of claim 19, wherein the composition has a freezing point of below -10°F.
- 10 34. The method of claim 19, wherein the composition has a freezing point of below -20°F.
35. The method of claim 19, wherein the freezing point depressant component makes up greater than 65 wt. % of the total composition.
- 15 36. The method of claim 19, wherein the freezing point depressant component makes up greater than 70 wt. % of the total composition.
37. The method of claim 19, wherein the freezing point depressant component  
20 makes up greater than 75 wt. % of the total composition.

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(54) Title: **ANTIMICROBIAL COMPOSITIONS FORMULATED FOR USE IN COLD TEMPERATURE CONDITIONS AND  
METHODS OF USE THEREOF**

(57) Abstract: Antimicrobial compositions containing a carboxylic acid, for example, a fatty acid, and a carrier medium including a freezing point depressant are disclosed. The compositions can be formulated for use as a teat dip, for use on milk producing animals. In one particularly advantageous embodiment, a composition is formulated as a teat dip and includes suitable emollients, skin conditioners and lubricants.

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 A61K9/00 A61K9/12 A61K47/10

According to International Patent Classification (IPC) or to both national classification and IPC

## B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

IPC 7 A01N A61K

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

EPO-Internal, WPI Data, PAJ, BIOSIS, MEDLINE, CHEM ABS Data, CAB Data

## C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	WO 00 13506 A (ALCIDE CORP ;KROSS ROBERT D (US)) 16 March 2000 (2000-03-16)	1,2, 4-16, 19-24, 26-37
Y	page 2, line 17 - line 19 page 1 -page 5, line 3 page 7	3,17,25
X	EP 0 906 724 A (ECOLAB INC) 7 April 1999 (1999-04-07)	18
Y	page 2, line 52 - line 58 page 5, line 27 - line 47; claims 1,6,20,24; examples IX,XIII,,XIV	3,17,25
A	WO 00 13507 A (ALCIDE CORP ;KROSS ROBERT D (US)) 16 March 2000 (2000-03-16) page 1 -page 4, line 18	1-37
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Further documents are listed in the continuation of box C.



Patent family members are listed in annex.

## \* Special categories of cited documents:

- \*A\* document defining the general state of the art which is not considered to be of particular relevance
- \*E\* earlier document but published on or after the international filing date
- \*L\* document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)
- \*O\* document referring to an oral disclosure, use, exhibition or other means
- \*P\* document published prior to the international filing date but later than the priority date claimed

\*T\* later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention

\*X\* document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone

\*Y\* document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.

\*Z\* document member of the same patent family

Date of the actual completion of the international search

20 June 2002

Date of mailing of the international search report

08/07/2002

Name and mailing address of the ISA

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## INTERNATIONAL SEARCH REPORT

International Application No

PC 01/30753

## C. (Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	DATABASE CA 'Online! CHEMICAL ABSTRACTS SERVICE, COLUMBUS, OHIO, US; WATANABE, HISAKI ET AL: "Antifreeze compositions" retrieved from STN Database accession no. 128:219346 CA XP002203008 abstract & JP 10 046134 A (NIPPON CHEMICAL INDUSTRIAL CO., LTD, JAPAN) 17 February 1998 (1998-02-17) -& PATENT ABSTRACTS OF JAPAN vol. 1998, no. 06, 30 April 1998 (1998-04-30) & JP 10 046134 A abstract	1-5, 7-16, 18
X	EP 0 864 627 A (CCI CO LIMITED) 16 September 1998 (1998-09-16) page 2, line 46 - line 58; claim 1; table 1	1-5, 7-16, 18
X	DD 231 065 A (LEIPZIG CHEMIEANLAGEN) 18 December 1985 (1985-12-18) example 2	1, 2, 4, 5, 7-16

# INTERNATIONAL SEARCH REPORT

International application No.  
PCT/US 01/30753

## Box I Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)

This International Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:

1. ☒ Claims Nos.:  
because they relate to subject matter not required to be searched by this Authority, namely:  
  
Although claims 19-37 are directed to a method of treatment of the animal body, the search has been carried out and based on the alleged effects of the composition.
2. ☐ Claims Nos.:  
because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:
3. ☐ Claims Nos.:  
because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).

## Box II Observations where unity of invention is lacking (Continuation of item 2 of first sheet)

This International Searching Authority found multiple inventions in this international application, as follows:

1. ☐ As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2. ☐ As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. ☐ As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4. ☐ No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:

### Remark on Protest

- ☐ The additional search fees were accompanied by the applicant's protest.
- ☐ No protest accompanied the payment of additional search fees.

## INTERNATIONAL SEARCH REPORT

International Application No

PC 01/30753

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